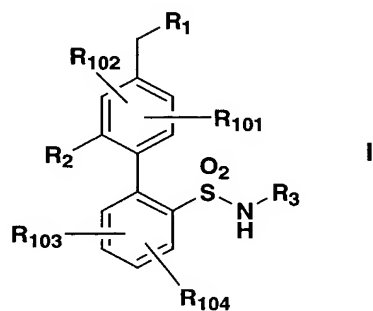


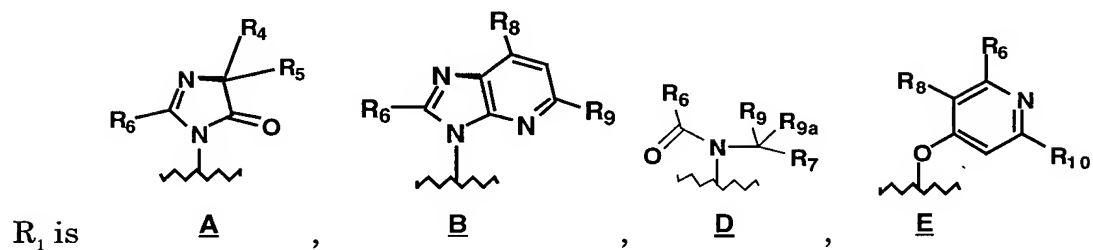
What is claimed is:

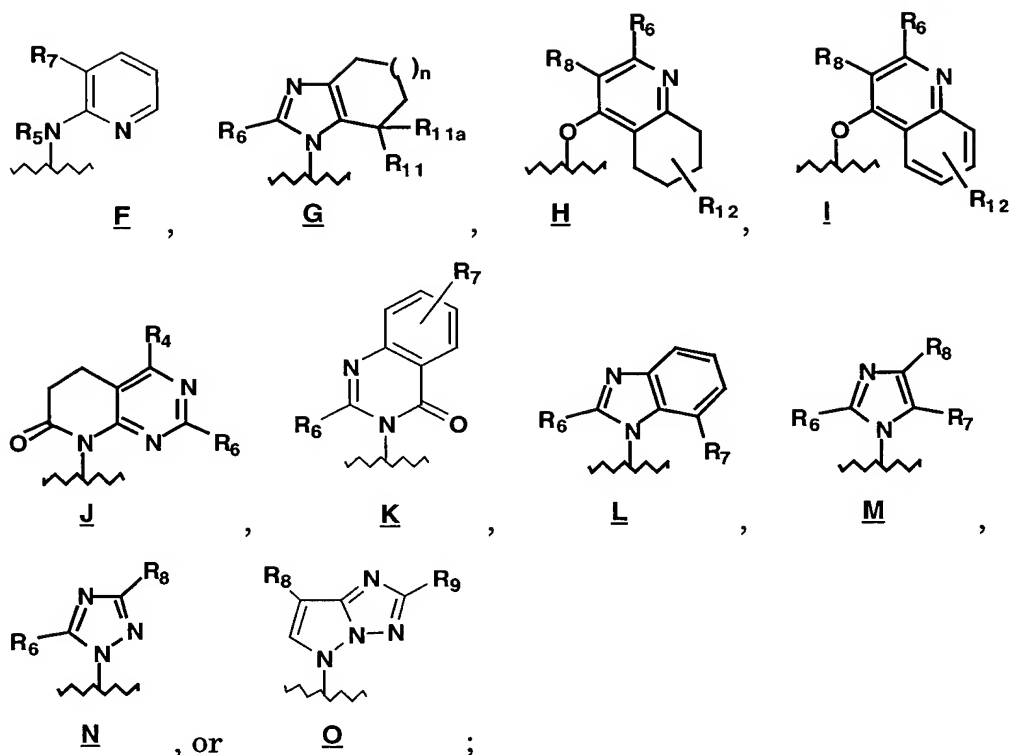
1. A compound of the following formula I, or an enantiomer, diastereomer, salt or metabolite thereof:

5



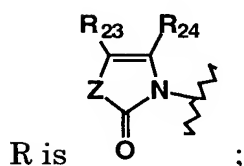
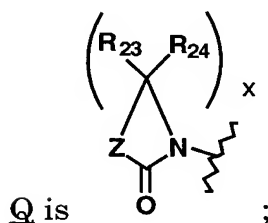
wherein:

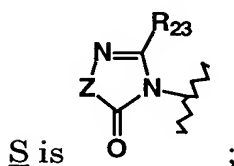




- 5  $R_2$  is hydrogen, halogen, -CHO, alkyl, haloalkyl, (cycloalkyl)alkyl, alkenyl, alkynyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, aryloxy alkoxyalkoxy, cyano, hydroxy, hydroxyalkyl, nitro,  $-\text{CH}(\text{OR}_{13})(\text{OR}_{14})$ ,  $-(\text{CH}_2)_w\text{Y}$ ; with the proviso that when  $R_1$  is B,  $R_2$  is not hydrogen, halogen, alkyl, haloalkyl, alkoxy, hydroxyalkyl, nitro,  $-(\text{CH}_2)_w\text{NR}_{19}\text{R}_{20}$  or  $-\text{NHSO}_2\text{R}_{22}$ ;
- 10  $R_3$  is heteroaryl;
- $R_4$  and  $R_5$  are each independently alkyl, hydroxyalkyl, cycloalkyl, hydroxy substituted cycloalkyl, alkoxyalkyl, or hydroxy substituted alkoxyalkyl, or  $R_4$  and  $R_5$  together form a cyclobutyl, cyclopentyl, cyclohexyl, tetrahydrofuranyl or tetrahydropyranyl ring which may
- 15 be optionally substituted with one or more hydroxy groups;
- $R_6$  is alkyl, hydroxyalkyl, haloalkyl, hydroxy substituted haloalkyl, cycloalkyl, hydroxy substituted cycloalkyl, (cycloalkyl)alkyl, hydroxy substituted (cycloalkyl)alkyl, aralkyl, alkoxy, hydroxy substituted alkoxy, alkoxyalkyl, hydroxy substituted alkoxyalkyl, or
- 20  $-\text{NR}_{16}\text{R}_{17}$ ;

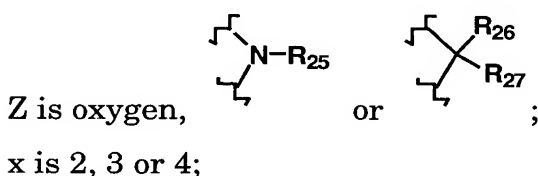
- $R_7$  is  $-(CH_2)_w-CO_2R_{15}$ ,  $-(CH_2)_w-(C=O)NR_{16}R_{17}$ ,  $-(CH_2)_w-NR_{15}(C=O)NR_{16}R_{17}$ ,  $-(CH_2)_w-CH_2OH$ ,  $-(CH_2)_w-(C=O)R_{15}$ , tetrazolyl, oxadiazolyl or triazolyl wherein said tetrazolyl, oxadiazolyl or triazolyl may optionally be substituted with hydrogen, alkyl, hydroxy or halogen;
- 5  $R_8$ ,  $R_9$ ,  $R_{9a}$ ,  $R_{10}$  and  $R_{12}$  are each independently hydrogen, halogen, alkyl, hydroxyalkyl, cycloalkyl, (cycloalkyl)alkyl, aryl, heteroaryl, arylalkyl, alkylthioalkyl, alkoxy or alkoxyalkyl, or  $R_9$  and  $R_{9a}$  together with the carbon atom to which they are bonded form a cycloalkyl ring;
- 10  $R_{11}$  and  $R_{11a}$  are each independently hydrogen, alkoxy, or together form a carbonyl;
- $R_{13}$  and  $R_{14}$  are alkyl or together form a five to six-membered ring;
- $R_{15}$ ,  $R_{16}$  and  $R_{17}$  are independently hydrogen, alkyl, hydroxyalkyl, cycloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, aralkyl, heterocycloalkyl,
- 15 aryl, heteroaryl or  $-(CH_2)_wQ$ , or  $R_{16}$  and  $R_{17}$  may together form a four to six-membered heterocyclic ring;
- $n$  is 1 or 2;
- $w$  is 0, 1, or 2;
- $Y$  is heteroaryl,  $-COOH$ ,  $-COOR_{18}$ ,  $-CONR_{19}R_{20}$ ,  $-NR_{19}R_{20}$ ,  $-NR_{19}-OR_{20}$ ,  $-NR_{21}(C=O)R_{22}$ ,  $-NR_{21}(C=O)NR_{19}R_{20}$ ,  $-N(R_{19})-(alk)-NR_{21}(C=O)R_{22}$ ,
- 20  $-NR_{21}(C=O)OR_{18}$ ,  $-NR_{21}SO_2R_{22}$ ,  $-SO_2R_{22}$ , Q, R or S;





$R_{18}$ ,  $R_{19}$ ,  $R_{20}$ ,  $R_{21}$  and  $R_{22}$  are each independently hydrogen, alkyl, haloalkyl, alkoxyalkyl, cycloalkyl, alkenyl, alkynyl, aryl, aralkyl, heteroaryl, or  $R_{19}$  and  $R_{20}$  may together form a four to seven-membered heterocyclic ring;

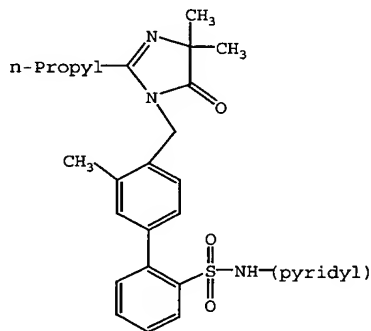
$R_{23}$  and  $R_{24}$  are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;



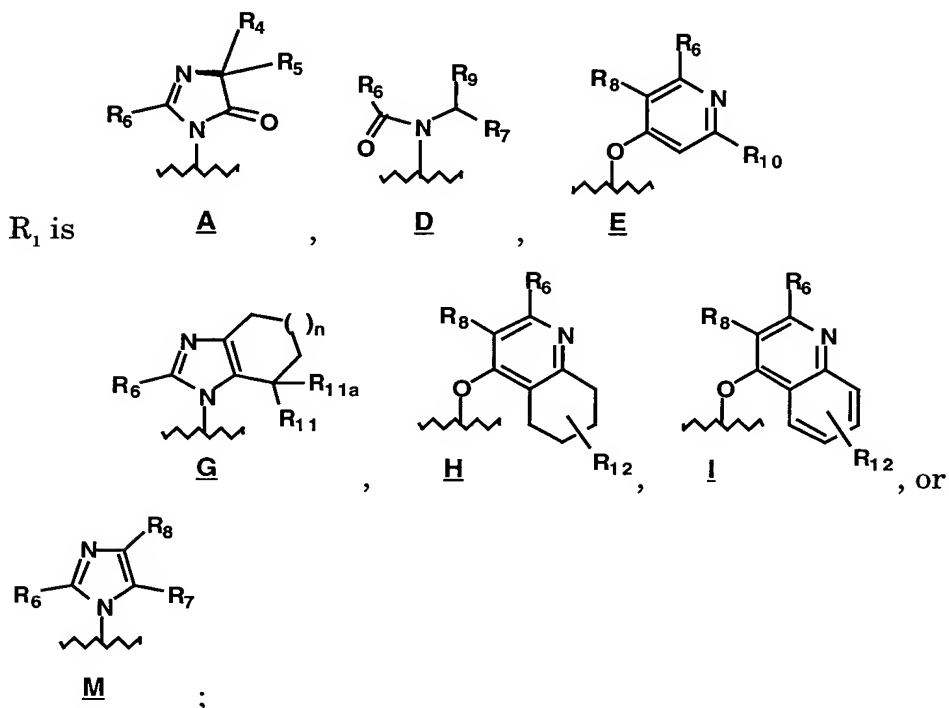
$R_{25}$ ,  $R_{26}$  and  $R_{27}$  are each independently hydrogen, alkyl or cycloalkyl, or  $R_{26}$  and  $R_{27}$  may together form a three to seven-membered cycloalkyl ring;

$R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are each independently hydrogen, halogen, -CHO, alkyl, haloalkyl, (cycloalkyl)alkyl, alkenyl, alkynyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, cyano, hydroxy, hydroxyalkyl, nitro, -CH(OR<sub>13</sub>)(OR<sub>14</sub>), or -(CH<sub>2</sub>)<sub>w</sub>Y; wherein said rings; aryl alone or as part of another group; or heteroaryl alone or as part of another group may each optionally be substituted by one or more hydrogen, halogen, cyano, alkyl, hydroxyalkyl, alkoxy, nitro or trifluoromethyl groups;

provided that when  $R_1$  is A said compound is other than



2. A compound of claim 1, wherein



$R_2$  is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or  $-(CH_2)_wY$ , or when  $R_1$  is D,  $R_2$  is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or  $-(CH_2)_wY$ ;

$R_3$  is isoxazolyl, pyridizynyl, pyrazinyl or pyrimidinyl, each optionally independently substituted with one to three substituents selected from hydrogen, halogen, cyano, alkyl, alkoxy, trifluoromethyl or nitro;

15  $R_4$  and  $R_5$  are each independently alkyl, cycloalkyl, or  $R_4$  and  $R_5$  together form a cyclobutyl, cyclopentyl or cyclohexyl ring;

$R_6$  is alkyl, haloalkyl, cycloalkyl or alkoxy;

$R_7$  is  $-CO_2R_{15}$ ,  $-(C=O)NR_{16}R_{17}$  or  $-CH_2OH$ ;

20  $R_8$ ,  $R_9$ ,  $R_{10}$  and  $R_{12}$  are each independently hydrogen, halogen, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;

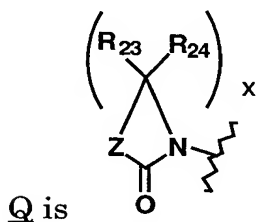
$R_{11}$  and  $R_{11a}$  are each independently hydrogen, alkoxy, or together form a carbonyl;

$R_{15}$ ,  $R_{16}$  and  $R_{17}$  are independently hydrogen, alkyl or cycloalkyl or  $R_{16}$  and  $R_{17}$  may together form a four to six-membered heterocyclic ring;

5  $n$  is 1 or 2;

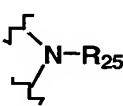
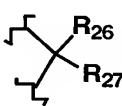
$w$  is 0, 1, or 2;

$Y$  is  $-\text{COOR}_{18}$ ,  $-\text{NR}_{21}(\text{C}=\text{O})\text{R}_{22}$ ,  $-\text{NR}_{21}(\text{C}=\text{O})\text{NR}_{19}\text{R}_{20}$ ,  $-\text{NR}_{21}(\text{C}=\text{O})\text{OR}_{18}$ ,  $-\text{NR}_{21}\text{SO}_2\text{R}_{22}$ ,  $-\text{SO}_2\text{R}_{22}$  or  $\underline{Q}$ ;



10  $R_{18}$ ,  $R_{19}$ ,  $R_{20}$ ,  $R_{21}$  and  $R_{22}$  are each independently hydrogen, alkyl, cycloalkyl, or  $R_{19}$  and  $R_{20}$  may together form a four to seven-membered heterocyclic ring;

$R_{23}$  and  $R_{24}$  are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;

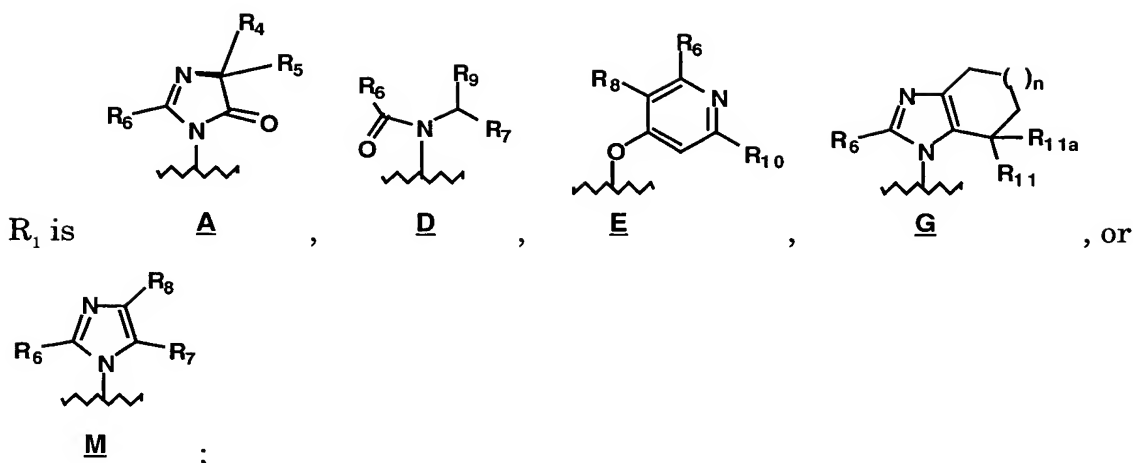
15  $Z$  is oxygen,  or  ;

$x$  is 2, 3 or 4;

$R_{25}$ ,  $R_{26}$  and  $R_{27}$  are each independently hydrogen, alkyl or cycloalkyl, or  $R_{26}$  and  $R_{27}$  may together form a three to seven-membered cycloalkyl ring;

20  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are each independently hydrogen, halogen, alkoxy or alkyl.

3. A compound of claim 1, wherein



$R_2$  is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, hydroxyalkyl, or  $-(CH_2)_wY$ , or when  $R_1$  is D,  $R_2$  is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or  $-(CH_2)_wY$ ;

$R_3$  is isoxazolyl, optionally independently substituted with one or two substituents selected from hydrogen, halogen, cyano, alkyl, alkoxy, trifluoromethyl or nitro;

$R_4$  and  $R_5$  are each independently alkyl, cycloalkyl, or  $R_4$  and  $R_5$  together form a cyclobutyl, cyclopentyl or cyclohexyl ring;

$R_6$  is alkyl, haloalkyl, cycloalkyl or alkoxy;

$R_7$  is  $-CO_2R_{15}$  or  $-(C=O)NR_{16}R_{17}$ ;

$R_8$ ,  $R_9$  and  $R_{10}$  are each independently hydrogen, halogen, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;

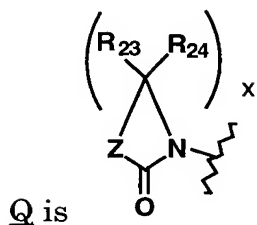
$R_{11}$  and  $R_{11a}$  together form a carbonyl;

$R_{15}$ ,  $R_{16}$  and  $R_{17}$  are independently hydrogen, alkyl, or cycloalkyl or  $R_{16}$  and  $R_{17}$  may together form a four to six-membered heterocyclic ring;

$n$  is 2;

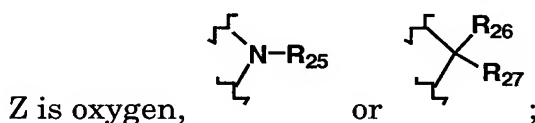
$w$  is 0, 1, or 2;

$Y$  is  $-NR_{21}(C=O)R_{22}$ ,  $-NR_{21}(C=O)NR_{19}R_{20}$ ,  $-NR_{21}(C=O)OR_{18}$ ,  $-NR_{21}SO_2R_{22}$ ,  $-SO_2R_{22}$  or Q;



$\text{R}_{18}$ ,  $\text{R}_{19}$ ,  $\text{R}_{20}$ ,  $\text{R}_{21}$  and  $\text{R}_{22}$  are each independently hydrogen, alkyl, cycloalkyl, or  $\text{R}_{19}$  and  $\text{R}_{20}$  may together form a four to seven-membered heterocyclic ring;

- 5  $\text{R}_{23}$  and  $\text{R}_{24}$  are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;

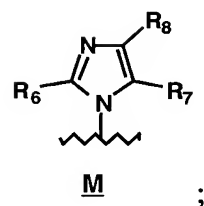
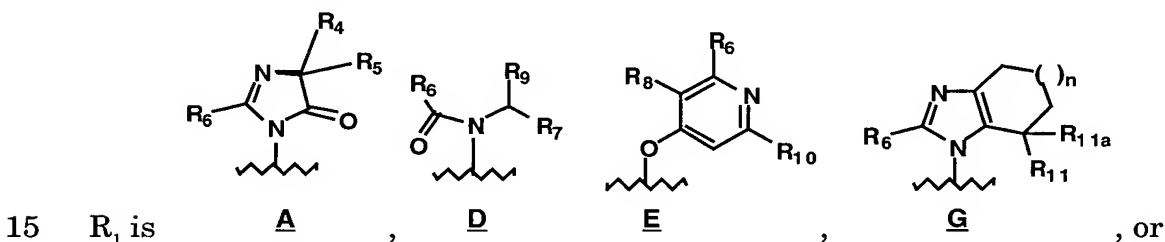


$x$  is 2, 3 or 4;

- 10  $\text{R}_{25}$ ,  $\text{R}_{26}$  and  $\text{R}_{27}$  are each independently hydrogen, alkyl or cycloalkyl, or  $\text{R}_{26}$  and  $\text{R}_{27}$  may together form a three to seven-membered cycloalkyl ring;

$\text{R}_{101}$ ,  $\text{R}_{102}$ ,  $\text{R}_{103}$ , and  $\text{R}_{104}$  are each independently hydrogen, halogen, or alkyl.

4. A compound of claim 1, wherein



$\text{R}_2$  is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or  $-(\text{CH}_2)_w\text{Y}$ ; or when  $\text{R}_1$  is  $\underline{\text{D}}$ ,  $\text{R}_2$



is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or  $-(CH_2)_wY$ ;

$R_3$  is isoxazol-5-yl or isoxazol-3-yl independently substituted with two

5 substituents selected from alkyl or halogen;

$R_4$  and  $R_5$  are each independently alkyl, cycloalkyl, or  $R_4$  and  $R_5$  together form a cyclobutyl, cyclopentyl or cyclohexyl ring;

$R_6$  is alkyl, haloalkyl, cycloalkyl or alkoxy;

$R_7$  is  $-(C=O)NR_{16}R_{17}$ ;

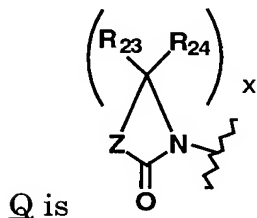
10  $R_8$ ,  $R_9$ , and  $R_{10}$  are independently H, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;

$n$  is 2;

$w$  is 0, 1, or 2;

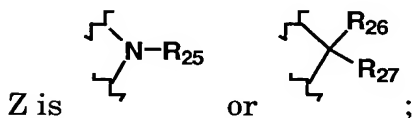
$Y$  is  $-NR_{21}(C=O)R_{22}$ ,  $-NR_{21}(C=O)NR_{19}R_{20}$ ,  $-NR_{21}(C=O)OR_{18}$ ,  $-NR_{21}SO_2R_{22}$

15 or  $Q$ ;



$R_{18}$ ,  $R_{19}$ ,  $R_{20}$ ,  $R_{21}$  and  $R_{22}$  are each independently hydrogen, alkyl, cycloalkyl, or  $R_{19}$  and  $R_{20}$  may together form a four-, five-, six- or to seven-membered heterocyclic ring;

20  $R_{23}$  and  $R_{24}$  are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;



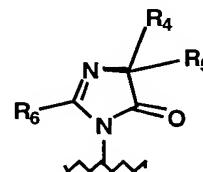
$x$  is 2;

$R_{25}$ ,  $R_{26}$  and  $R_{27}$  are each independently hydrogen, alkyl or cycloalkyl, or  $R_{26}$

25 and  $R_{27}$  may together form a three-, four-, five, six- or seven-membered cycloalkyl ring;

$R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are each independently hydrogen, halogen, or alkyl.

5. A compound of claim 1, wherein  $R_3$  is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.

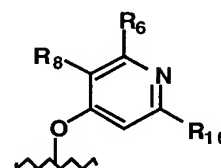


6. A compound of claim 5, wherein  $R_1$  is A.

7. A compound of claim 6, wherein  $R_2$  is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ ,  $R_{104}$  are each independently hydrogen, halogen, or alkyl.

8. A compound of claim 6, wherein  $R_2$  is  $-CH_2Y$ .

9. A compound of claim 8, wherein Y is Q.

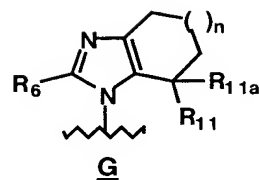


10. A compound of claim 5, wherein  $R_1$  is E.

11. A compound of claim 10, wherein  $R_2$  is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ ,  $R_{104}$  are each independently hydrogen, halogen, or alkyl.

12. A compound of claim 10, wherein  $R_2$  is  $-CH_2Y$ .

13. A compound of claim 12, wherein Y is Q.



14. A compound of claim 5, wherein  $R_1$  is G.

15. A compound of claim 14, wherein  $R_2$  is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ ,  $R_{104}$  are each independently hydrogen, halogen, or alkyl.

16. A compound of claim 14, wherein  $R_2$  is  $-CH_2Y$ .

17. A compound of claim 16, wherein Y is Q.

18. A compound of claim 1, wherein  $R_2$  is alkoxyalkyl alkyl, haloalkyl or haloalkoxyalkyl

19. A compound of claim 18, wherein  $R_3$  is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.

20. A compound of claim 1, wherein  $R_2$  is  $-CH_2Y$ .

21. A compound of claim 20, wherein  $R_3$  is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.

22. A compound of claim 20, wherein Y is Q.

23. A compound of claim 22, wherein R<sub>3</sub> is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.

5 24. A compound of claim 1, wherein said compound is selected from the group consisting of

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylamino)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3-methyl-2-oxo-1-imidazolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(2-pyrazinyl) [1,1'-biphenyl]-2-sulfonamide;

25 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3-chloro-2-pyrazinyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(3,6-dimethyl-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(3-methoxy-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- (S)-N-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-[2-methyl-1-(3-methyl-1,2,4-oxadiazol-5-yl)propyl]pentanamide;
- 15 N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[2-(2-methoxyethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 20 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[2-(ethoxymethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[(2-oxo-1-pyrrolidinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 25 4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[(3-methyl-2-oxo-1-imidazolidinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- (S)-2-[N-[2'-[[N-(3-Methyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;
- 30 (S)-2-[N-[2'-[[N-(4-Bromo-3-methyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]  
-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(7-Methoxycarbonyl-2-ethoxybenzimidazol-1-yl)methyl]-  
N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(7-Carboxy-2-ethoxybenzimidazol-1-yl)methyl]-N-  
(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(7-Methoxycarbonyl-2-ethylbenzimidazol-1-yl)methyl]-N-  
(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(7-Carboxy-2-ethylbenzimidazol-1-yl)methyl]-N-  
10 (3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-  
(methoxycarbonyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-  
isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-  
15 (carboxy)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-  
biphenyl]-2-sulfonamide;
- 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N-  
methylcarbamoyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-  
[1,1'-biphenyl]-2-sulfonamide;
- 20 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N,N-  
dimethylcarbamoyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-  
isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(1,3,5-trimethylpyrazol-4-yl)  
[1,1'-biphenyl]-2-sulfonamide;
- 25 4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(3-methylisoxazol-5-yl) [1,1'-  
biphenyl]-2-sulfonamide;
- 4'-[(5-Acetyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(3,4-  
dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(5-Methoxycarbonyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-  
30 (3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

- 4'-[(5-(N,N-dimethylcarbamoyl)-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-hydroxymethyl[1,1'-biphenyl]-2-sulfonamide
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidiny)methyl]-N-(3-methoxy-5-methyl-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-bromo-3-methyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidiny)methyl][1,1'-biphenyl]-2-sulfonamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(formylmethylamino)-methyl][1,1'-biphenyl]-2-sulfonamide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylpropanamide;
- 20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylcyclopropanecarbox-amide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,2-dimethylpropanamide;
- 25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
 methoxy-N-methylacetamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methyl-4-pentynamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methylcyclobutanecarboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,3-dimethylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 15 N,2,2-trimethylpropanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
 methoxy-N-methylpropanamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
 ethoxy-N-methylacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methyl-2-furancarboxamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,4-dimethylpentanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 30 methylbenzamide;



N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
methyl-3-thiophenecarboxamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
methylcyclopentaneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
methylcyclohexanecarboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
N,3-dimethylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
15 methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
20 [[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-  
fluoro-N-methylbenzamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
methylcyclohexaneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
30 fluoro-N-methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
 fluoro-N-methylbenzeneacetamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-  
 fluoro-N-methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,N',N'-trimethylurea;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N'-  
 (1,1-dimethylethyl)-N-methylurea;

15 [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-  
 yl)methyl)methylcarbamic acid ethyl ester;

[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-  
 yl)methyl)methylcarbamic acid 2-methylpropyl ester;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,3,3-trimethylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methyl-2-pyridinecarboxamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methyl-3-pyridinecarboxamide;

30 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methyl-2-pyrazinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,1-dimethyl-1H-pyrrole-2-carboxamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methyl-1,2,3-thiadiazole-4-carboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,5-dimethyl-2-pyrazinecarboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,3,5-trimethyl-4-isoxazolecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 15 N,3-dimethyl-2-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,5-dimethyl-2-thiophenecarboxamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
 cyano-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-  
 cyano-N-methylbenzamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
 methoxy-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
 30 chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
 chloro-N-methylbenzamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-  
 chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 2,3-difluoro-N-methylbenzamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 3,4-difluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 15 3,5-difluoro-N-methylbenzamide;

4-Acetyl-N-[[4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-  
 yl)methyl]-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-  
 yl)methyl]-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 20 [(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
 ethoxy-N-methyl-2-thiophenecarboxamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-  
 dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-  
 25 dimethyl-5-isoxazolyl)-2'-[(propylsulfonyl)amino][1,1'-biphenyl]-2-  
 sulfonamide;

N-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-  
 yl)methyl]-N-(1-oxopentyl)-L-valine methyl ester;

N-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-  
 30 yl)methyl]-N-(1-oxopentyl)-L-valine;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidiny)methyl]-4'-[(2-propyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

5 N-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl]methyl]-N,3,3-trimethylbutanamide;

N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N2-(1-oxopentyl)-L-valinamide;

10 N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxopentyl)-L-valinamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2,2,2-trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid methyl ester;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

25 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;

2'-[Cyanomethyl]-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

30 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(cyanomethyl)-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-cyano-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 5 2'-Cyano-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-methyl[1,1'-biphenyl]-4-yl]methyl-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[(2,2,2-trifluoroethyl)amino]methyl[1,1'-biphenyl]-2-sulfonamide;
- 10 N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]benzeneacetamide;
- 15 N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]-3,3-dimethylbutanamide;
- 2'-Amino-4'-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-nitro[1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl-N-methyl-N2-(1-oxopropyl)-L-isoleucinamide;
- N2-(Cyclopropylcarbonyl)-N2-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl-N-methyl-L-isoleucinamide;
- 25 N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl-N-methyl-N2-(1-oxo-3-phenylpropyl)-L-isoleucinamide;
- N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl-N-methyl-N2-(3-methyl-1-oxobutyl)-L-isoleucinamide;
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- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-isoleucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-isoleucinamide;
- 5 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-leucinamide;
- N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-leucinamide;
- 10 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-leucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(phenylacetyl)-L-leucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-leucinamide;
- 15 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-leucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-leucinamide;
- 20 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-valinamide;
- N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-valinamide;
- 25 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(phenylacetyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-valinamide;
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- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-valinamide;
- 5 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-ethyl- N2-(1-oxopentyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-hexyl- N2-(1-oxopentyl)-L-valinamide;
- 10 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-cyano[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-hydroxymethyl[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 15 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- N,N-Dimethyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- 20 N-Methyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- 25 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;



N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

5 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

2'-Chloro-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[[(5,6,7,8-tetrahydro-2-ethyl-4-quinolinyloxy)methyl][1,1'-biphenyl]-2-sulfonamide;

10 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-fluoro[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(phenoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

15 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-(Cyclopropylcarbonyl)-N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

20 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

25 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

N2-[[2-Chloro-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

30 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-(Cyclobutylcarbonyl)-N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-L-valinamide;

1-[[2-Chloro-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxylic acid;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

(S)-N-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-[2-methyl-1-[(4-methyl-1-piperazinyl)carbonyl]propyl]pentanamide;

(S)-N-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-piperidiny)carbonyl]propyl]pentanamide;

N-(3,3-Dimethylbutyl)- N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-[(4-fluorophenyl)methyl]-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(propoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4-Chloro-1-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-propyl-1H-imidazole-5-carboxamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(1,2-dihydro-2-oxo-1-pyridinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-2-sulfonamide;
- 2-Butyl-4-chloro-1-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-1H-imidazole-5-carboxamide;
- 5 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2-methyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2-ethyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- 10 [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2-ethyl-5,6,7,8-tetrahydro-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2-propyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- 15 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(5,6,7,8-tetrahydro-2,4-dimethyl-7-oxopyrido[2,3-d]pyrimidin-8-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[[2-Ethyl-4-quinolinyloxy]methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 20 N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[[5,6,7,8-tetrahydro-2-ethyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- 3-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-N-methyl-3H-benzimidazole-4-carboxamide;
- 25 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid phenylmethyl ester;
- 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-phenylethyl ester;
- 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-(2-oxo-1-pyrrolidinyl)ethyl ester;
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- 1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 3-(2-oxo-1-pyrrolidinyl)propyl ester;
- 2'-Cyano-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[[(2-ethyl-4-quinolinyloxy)methyl][1,1'-biphenyl]-2-sulfonamide;
- 2'-(Cyanomethyl)-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 3-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-5-ethyl-N-methyl-2-propyl-3H-imidazole-4-carboxamide;
- 1-[[[2-Chloro-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 3-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-5-ethyl-2-propyl-3H-imidazole-4-carboxamide;
- 3-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethoxy-N-methyl-3H-benzimidazole-4-carboxamide;
- 3-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethoxy-N,N-dimethyl-3H-benzimidazole-4-carboxamide;
- 2-[[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]propylamino]-3-pyridinecarboxylic acid;
- 4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2,7-Diethyl-5H-pyrazolo[1,5-b][1,2,4]triazol-5-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- N-[2-Butyl-3-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-3,4-dihydro-4-oxo-6-quinazolinyl]-N'-methyl-N'-(1-methylethyl)urea;
- 2-[[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]propylamino]-N-methyl-3-pyridinecarboxamide;

- 4'-[[[(3-Methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- 5 N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- 10 3-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethyl-5-methyl-3H-imidazole-4-carboxamide;
- 4'-[(2-Butyl-3,4-dihydro-4-oxo-3-quinazolinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(4,4-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 25 N-[2-[[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl]methyl]methylamino]ethyl]acetamide;

4-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-  
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-acetic acid  
ethyl ester;

5 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-  
4-yl)methyl]-N2-(1-oxopentyl)-N-propyl-L-valinamide;

N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-  
4-yl)methyl]-N2-(1-oxopentyl)-N-[(tetrahydro-2-furanyl)methyl]-L-  
valinamide;

10 2'-Chloro-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[[(2-ethyl-4-  
quinolinyloxy)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(2-ethyl-4-quinolinyloxy)methyl]-  
2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-chloro-  
N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-  
dimethyl-5-isoxazolyl)-2'-[(2-methylpropoxy)methyl][1,1'-biphenyl]-2-  
sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-  
dimethyl-5-isoxazolyl)-2'-[(ethylsulfonyl)amino][1,1'-biphenyl]-2-  
20 sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-  
dimethyl-5-isoxazolyl)-2'-[(2,2,2-trifluoroethoxy)methyl][1,1'-biphenyl]-2-  
sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-  
25 dimethyl-5-isoxazolyl)-2'-[(2-fluoroethoxy)methyl][1,1'-biphenyl]-2-  
sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-  
pyridinyl)oxy)methyl]-2'-ethoxymethyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-  
30 dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(3,3,3-trifluoropropyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(3-fluoropropyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(1,1-difluoroethyl)-N-(3,4-dimethylisoxazol-5-yl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2,2-trifluoroethyl)-N-(3,4-dimethylisoxazol-5-yl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-[(2-methyl)propoxy] [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(2-methoxyethoxy) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-N-(3,4-dimethylisoxazol-5-yl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3-methylisoxazol-5-yl)-2'-trifluoromethyl[1,1'-biphenyl]-2-sulfonamide;

N-(4-Bromo-3-methyl-5-isoxazolyl)-4'-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-trifluoromethyl [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-chloro-3-methyl-5-isoxazolyl)-2'-trifluoromethyl [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(N-methoxy-N-methylaminomethyl) [1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2-difluoroethoxymethyl)-N-(3,4-dimethylisoxazol-5-yl)[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(2-fluoroethyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(2-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(3-methylbutyl)[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(2-methylpropyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[[2-(3,3-Difluorobutyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- 15 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-[(1,1-dimethylethoxy)methyl][1,1'-biphenyl]-2-
- 20 sulfonamide;
- 1-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 1-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;
- 25 1-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 1-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-
- 30 carboxamide;



N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethoxymethyl-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl] [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2-fluoroethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl] [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl] [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethoxymethyl-4'-[[4-oxo-2-(3,3,3-trifluoropropyl)-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl] [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1-hydroxyethyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

1-[[2'-[[3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

- N,4-Diethyl-1-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-2-propyl-1H-imidazole-5-carboxamide;  
 1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;  
 5 1-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-(1-methylethyl)-2-propyl-1H-imidazole-5-carboxamide;  
 1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;  
 10 imidazole-5-carboxamide;  
 N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethoxymethyl-4'-[[[(6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;  
 N2-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxobutyl)-L-valinamide;  
 15 N2-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxopentyl)-L-valinamide;  
 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;  
 4'-[(5-Acetyl-4-ethyl-2-propylimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;  
 20 N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;  
 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;  
 25 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;  
 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;  
 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;  
 30

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-3'-fluoro-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-3'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-
- 20 sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-
- 25 N-(4,5-dimethyl-3-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4,5-dimethoxy[1,1'-biphenyl]-2-
- 30 sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-4-methoxy-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
- 10 dimethyl-3-isoxazolyl)-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-5-fluoro[1,1'-biphenyl]-2-sulfonamide;
- N<sup>2</sup>-(Cyclopropylcarbonyl)-N<sup>2</sup>-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
- 15 valinamide;
- N<sup>2</sup>-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,3-dimethyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
- N<sup>2</sup>-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N<sup>2</sup>-(2-methyl-1-oxopropyl)-L-valinamide;
- 20 N<sup>2</sup>-(Cyclopentylcarbonyl)-N<sup>2</sup>-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-valinamide;
- N<sup>2</sup>-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-3-fluoro[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
- 25 N<sup>2</sup>-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(1-methylethyl)-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
- N<sup>2</sup>-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(2-methoxyethyl)-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
- N-(Cyclopropylmethyl)-N<sup>2</sup>-[[2'-[(4,5-dimethyl-3-
- 30 isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

N<sup>2</sup>-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N<sup>2</sup>-(1-oxobutyl)-N-(3-pyridinyl)-L-valinamide;

N<sup>2</sup>-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-N<sup>2</sup>-(1-oxopentyl)-L-valinamide;

5 N-Methyl-N<sup>2</sup>-[[2'-[[[5-methyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N<sup>2</sup>-(1-oxopentyl)-L-valinamide;

N<sup>2</sup>-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-ethyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

10 N<sup>2</sup>-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-5'-fluoro[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[2,6-dimethyl-3-methoxy-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[2,6-dimethyl-3-methoxy-4-pyridinyl)oxy]methyl]-3'-fluoro[1,1'-biphenyl]-2-sulfonamide;

15 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-3'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

20 N-(4,5-Dimethyl-3-isoxazolyl)-5-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

25 N-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(ethoxymethyl)[1,1'-biphenyl]-4-yl)methyl]-1-[(1-oxopentyl)amino]cyclopentanamide;

and salts, enantiomers and diastereomer thereof.

25. A compound of claim 1, wherein said compound is selected from the group consisting of

30 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(methylamino)methyl][1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3-methyl-2-oxo-1-imidazolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 20 (S)-N-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-[2-methyl-1-(3-methyl-1,2,4-oxadiazol-5-yl)propyl]pentanamide;
- N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 25 N-(4,5-dimethyl-3-isoxazolyl)-4'-[2-(2-methoxyethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[2-(ethoxymethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-  
[(2-oxo-1-pyrrolidinyl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-  
2-sulfonamide;

5 4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-  
[(3-methyl-2-oxo-1-imidazolidinyl)methyl]-N-(4,5-dimethyl-3-  
isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

(S)-2-[N-[2'-[[N-(5-Methyl-3-isoxazolyl)amino]sulfonyl][1,1'-  
biphenyl]-4-yl)methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;

10 (S)-2-[N-[2'-[[N-(4-Bromo-5-methyl-3-isoxazolyl)amino]sulfonyl][1,1'-  
biphenyl]-4-yl)methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]  
-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

4'-[(7-Methoxycarbonyl-2-ethoxybenzimidazol-1-yl)methyl]-  
N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(7-Carboxy-2-ethoxybenzimidazol-1-yl)methyl]-N-  
(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

4'-[(7-Methoxycarbonyl-2-ethylbenzimidazol-1-yl)methyl]-N-  
(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

20 4'-[(7-Carboxy-2-ethylbenzimidazol-1-yl)methyl]-N-  
(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-  
(methoxycarbonyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-  
isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

25 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-  
(carboxy)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-  
biphenyl]-2-sulfonamide;

2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N-  
methylcarbamoyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-  
[1,1'-biphenyl]-2-sulfonamide;

- 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N,N-dimethylcarbamoyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(5-methylisoxazol-3-yl)[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(5-Acetyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(5-Methoxycarbonyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(5-(N,N-dimethylcarbamoyl)-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-hydroxymethyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-bromo-5-methyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(formylmethylamino)-methyl][1,1'-biphenyl]-2-sulfonamide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-methylpropanamide;
- 25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-methylcyclopropanecarbox-amide;



N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,2-dimethylpropanamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
 methoxy-N-methylacetamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methyl-4-pentynamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 15 methylcyclobutanecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,3-dimethylbutanamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,2,2-trimethylpropanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
 methoxy-N-methylpropanamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
 ethoxy-N-methylacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 30 methyl-2-furancarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
 N,4-dimethylpentanamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methyl-3-thiophenecarboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methylcyclopentaneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 15 methylcyclohexanecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 N,3-dimethylbenzamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
 methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-  
 fluoro-N-methylbenzamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
 fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-  
 30 fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-methylcyclohexaneacetamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-fluoro-N-methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-fluoro-N-methylbenzeneacetamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-fluoro-N-methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N,N',N'-trimethylurea;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N'-(1,1-dimethylethyl)-N-methylurea;

20 [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]methylcarbamic acid ethyl ester;

[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]methylcarbamic acid 2-methylpropyl ester;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N,3,3-trimethylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-methyl-2-pyridinecarboxamide;

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N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
methyl-3-pyridinecarboxamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
methyl-2-pyrazinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
N,1-dimethyl-1H-pyrrole-2-carboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-  
methyl-1,2,3-thiadiazole-4-carboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
15 N,5-dimethyl-2-pyrazinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
N,3,5-trimethyl-4-isoxazolecarboxamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
N,3-dimethyl-2-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-  
N,5-dimethyl-2-thiophenecarboxamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-  
cyano-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-  
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-  
30 cyano-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-methoxy-N-methylbenzamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-chloro-N-methylbenzamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2,3-difluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3,4-difluoro-N-methylbenzamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3,5-difluoro-N-methylbenzamide;

4-Acetyl-N-[[4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-methylbenzamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-ethoxy-N-methyl-2-thiophenecarboxamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(propylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

5 N-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-oxopentyl)-L-valine methyl ester;

N-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-oxopentyl)-L-valine;

10 N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(2-propyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl)methyl]-N,3,3-trimethylbutanamide;

15 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[[[(2,2,2-trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;

4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

25 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid methyl ester;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

30 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;

- 2'-(Cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-cyano-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 2'-Cyano-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-methyl[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[[[(2,2,2-trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;
- N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]benzeneacetamide;
- N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]-3,3-dimethylbutanamide;
- 2'-Amino-4'-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-nitro[1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopropyl)-L-isoleucinamide;
- N2-(Cyclopropylcarbonyl)-N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-isoleucinamide;

- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-isoleucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-isoleucinamide;
- 5 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxohexyl)-L-isoleucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxobutyl)-L-isoleucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxopropyl)-L-leucinamide;
- 10 N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-L-leucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-leucinamide;
- 15 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(phenylacetyl)-L-leucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-leucinamide;
- 20 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxohexyl)-L-leucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxobutyl)-L-leucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxopropyl)-L-valinamide;
- 25 N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-valinamide;
- 30



- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(phenylacetyl)-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-valinamide;
- 5 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxohexyl)-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxobutyl)-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-ethyl- N2-(1-oxopentyl)-L-valinamide;
- 10 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-hexyl- N2-(1-oxopentyl)-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-cyano[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 15 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-hydroxymethyl[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- 20 N,N-Dimethyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- N-Methyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- 25 N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 30

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

5 N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

10 2'-Chloro-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[5,6,7,8-tetrahydro-2-ethyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-fluoro[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(phenoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

20 N2-(Cyclopropylcarbonyl)-N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

25 N2-(Cyclopropylcarbonyl)-N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

30 N2-[[2-Chloro-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(trifluoromethyl)[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

5 N2-(Cyclobutylcarbonyl)-N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-L-valinamide;

1-[[2-Chloro-2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxylic acid;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-  
10 dimethyl-3-isoxazolyl)-2'-[(methylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

(S)-N-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-[2-methyl-1-[(4-methyl-1-piperazinyl)carbonyl]propyl]pentanamide;

15 (S)-N-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-piperidiny)carbonyl]propyl]pentanamide;

N-(3,3-Dimethylbutyl)- N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-L-valinamide;

20 N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-[(4-fluorophenyl)methyl]-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(methylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

25 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(propoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4-Chloro-1-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-propyl-1H-imidazole-5-carboxamide;

N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-  
30 oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(1,2-dihydro-2-oxo-1-pyridinyl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-2-sulfonamide;
- 2-Butyl-4-chloro-1-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-1H-imidazole-5-carboxamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-methyl-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,6,7,8-tetrahydro-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-propyl-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(5,6,7,8-tetrahydro-2,4-dimethyl-7-oxopyrido[2,3-d]pyrimidin-8-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Ethyl-4-quinolinyl)oxy]methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(5,6,7,8-tetrahydro-2-ethyl-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;
- 3-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-N-methyl-3H-benzimidazole-4-carboxamide;
- 1-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid phenylmethyl ester;
- 1-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-phenylethyl ester;

1-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-(2-oxo-1-pyrrolidinyl)ethyl ester;

5 1-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 3-(2-oxo-1-pyrrolidinyl)propyl ester;

2'-Cyano-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[[(2-ethyl-4-quinolinyloxy)methyl][1,1'-biphenyl]-2-sulfonamide;

10 2'-(Cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

3-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-5-ethyl-N-methyl-2-propyl-3H-imidazole-4-carboxamide;

15 1-[[2-Chloro-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

3-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-5-ethyl-2-propyl-3H-imidazole-4-carboxamide;

3-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethoxy-N-methyl-3H-benzimidazole-4-carboxamide;

20 3-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethoxy-N,N-dimethyl-3H-benzimidazole-4-carboxamide;

2-[[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]propylamino]-3-pyridinecarboxylic acid;

25 4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2,7-Diethyl-5H-pyrazolo[1,5-b][1,2,4]triazol-5-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

N-[2-Butyl-3-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-3,4-dihydro-4-oxo-6-quinazolinyl]-N'-methyl-N'-(1-methylethyl)urea;

2-[[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]propylamino]-N-methyl-3-pyridinecarboxamide;

4'-[[[3-Methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

3-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethyl-5-methyl-3H-imidazole-4-carboxamide;

4'-[(2-Butyl-3,4-dihydro-4-oxo-3-quinazolinyl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(4,4-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-[2-[[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl)methyl]methylamino]ethyl]acetamide;

4-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-acetic acid ethyl ester;

N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-N-propyl-L-valinamide;

N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-N-[(tetrahydro-2-furanyl)methyl]-L-valinamide;

2'-Chloro-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[[2-ethyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[[[2-ethyl-4-quinolinyloxy]methyl]-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-chloro-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-methylpropoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(ethylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2,2,2-trifluoroethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-methoxy-2,6-dimethyl-4-pyridinyloxy]methyl]-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(3,3,3-trifluoropropyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(3-fluoropropyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(1,1-difluoroethyl)-N-(4,5-dimethylisoxazol-3-yl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2,2-trifluoroethyl)-N-(4,5-dimethylisoxazol-3-yl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-[(2-methyl)propoxy] [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-methoxyethoxy) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-N-(4,5-dimethylisoxazol-3-yl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(N-methoxy-N-methylaminomethyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2-difluoroethoxymethyl)-N-(4,5-dimethylisoxazol-3-yl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-fluoroethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(3-methylbutyl)[1,1'-biphenyl]-2-sulfonamide;



- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-methypropyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[[2-(3,3-Difluorobutyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(ethoxymethyl)[1,1'-biphenyl]-
- 5 2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
- 10 dimethylisoxazol-3-yl)-2'-[(1,1-dimethylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;
- 1-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 15 1-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;
- 1-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 20 1-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;
- N,4-Diethyl-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl)methyl]-2-propyl-1H-imidazole-5-carboxamide;
- 25 1-[[2'-[[3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-(1-methylethyl)-2-propyl-1H-imidazole-5-carboxamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethoxymethyl-4'-[[6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- 30 4'-[(5-Acetyl-4-ethyl-2-propylimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy)methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
- 15 dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-fluoro-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
- 20 N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
- 25 dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4-methoxy-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-5-fluoro[1,1'-biphenyl]-2-sulfonamide;
- N<sup>2</sup>-(Cyclopropylcarbonyl)-N<sup>2</sup>-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
- 25 valinamide;
- N<sup>2</sup>-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,3-dimethyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
- N<sup>2</sup>-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N<sup>2</sup>-(2-methyl-1-oxopropyl)-L-valinamide;

N<sup>2</sup>-(Cyclopentylcarbonyl)-N<sup>2</sup>-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-L-valinamide;

5 N<sup>2</sup>-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-3-fluoro[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

N<sup>2</sup>-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-methylethyl)-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

N<sup>2</sup>-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(2-methoxyethyl)-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

10 N-(Cyclopropylmethyl)-N<sup>2</sup>-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

N<sup>2</sup>-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N<sup>2</sup>-(1-oxobutyl)-N-(3-pyridinyl)-L-valinamide;

15 N<sup>2</sup>-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-N<sup>2</sup>-(1-oxopentyl)-L-valinamide;

N-Methyl-N<sup>2</sup>-[[2'-[(3-methyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N<sup>2</sup>-(1-oxopentyl)-L-valinamide;

20 N<sup>2</sup>-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-ethyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

N<sup>2</sup>-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-5'-fluoro[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2,6-dimethyl-3-methoxy-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

25 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2,6-dimethyl-3-methoxy-4-pyridinyl)oxy]methyl]-3'-fluoro[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

30 N-(3,4-Dimethyl-5-isoxazolyl)-3'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-5-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

5 N-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(ethoxymethyl)[1,1'-biphenyl]-4-yl)methyl]-1-[(1-oxopentyl)amino]cyclopentanamide;

and salts, enantiomers and diastereomer thereof.

10 26. A method for the treatment of an endothelin-dependent or angiotensin II-dependent disorder, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

15 27. A method for the treatment of hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

20 28. A method for the treatment of pulmonary hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

25 29. A method for the treatment of primary pulmonary hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

30 30. A method for the treatment of low renin hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

31. A method for the treatment of male erectile dysfunction, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

5           32. A method for the treatment of male or female sexual dysfunction, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

10           33. A method for the treatment of heart failure, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

15           34. A method for the treatment of atherosclerosis, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

20           35. A method for the treatment of restenosis, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

            36. A method for the treatment of endotoxemia, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

25           37. A method for the inhibition of cell growth, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

30           38. A method for the treatment of cancer, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

39. A method for the treatment of migraine, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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40. A method for the treatment of asthma, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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41. A method for the treatment of ischemia, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

15

42. A method for the treatment of subarachnoid hemorrhage, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

20

43. A method for the treatment of benign prostatic hypertrophy, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

25

44. A method for the treatment of renal, glomerular or mesangial cell disorders, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

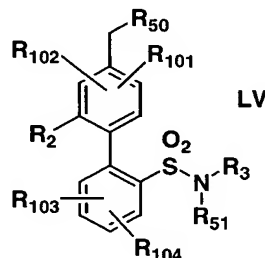
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45. A method for the treatment of acute or chronic renal failure, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

46. A pharmaceutical composition for the treatment of an endothelin-dependent or angiotensin II-dependent disorder, comprising a

pharmaceutically acceptable vehicle or diluent and at least one compound of claim 1 in an amount effective therefor.

47. A compound of the formula



5

wherein

$R_2$ ,  $R_3$ ,  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are as defined in claim 1;

$R_{50}$  is hydroxy, chloro, bromo, iodo,  $-\text{OSO}_2\text{-alkyl}$ , or  $-\text{OSO}_2\text{-aryl}$ ; and

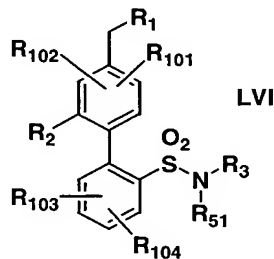
$R_{51}$  is hydrogen, or a suitable nitrogen protecting group.

10

48. The compound of claim 47, wherein  $R_{51}$  is

$-\text{CH}_2\text{OCH}_2\text{CH}_2\text{OCH}_3$ ,  $-\text{CH}_2\text{OCH}_2\text{CH}_2\text{Si}(\text{CH}_3)_3$ ,  $-\text{CH}_2\text{OCH}_3$ , or  $-\text{CH}_2\text{OCH}_2\text{-aryl}$ .

49. A compound of the formula



15

wherein  $R_1$ ,  $R_2$ ,  $R_3$ ,  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are as defined in claim 1; and

$R_{51}$  is a suitable nitrogen protecting group.

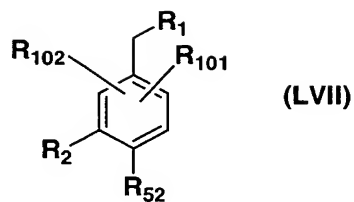
50. The compound of claim 49, wherein  $R_{51}$  is  $-\text{CH}_2\text{OCH}_2\text{CH}_2\text{OCH}_3$ , -

20

$\text{CH}_2\text{OCH}_2\text{CH}_2\text{Si}(\text{CH}_3)_3$ ,  $-\text{CH}_2\text{OCH}_3$ , or  $-\text{CH}_2\text{OCH}_2\text{-aryl}$ .

51. A compound of the formula





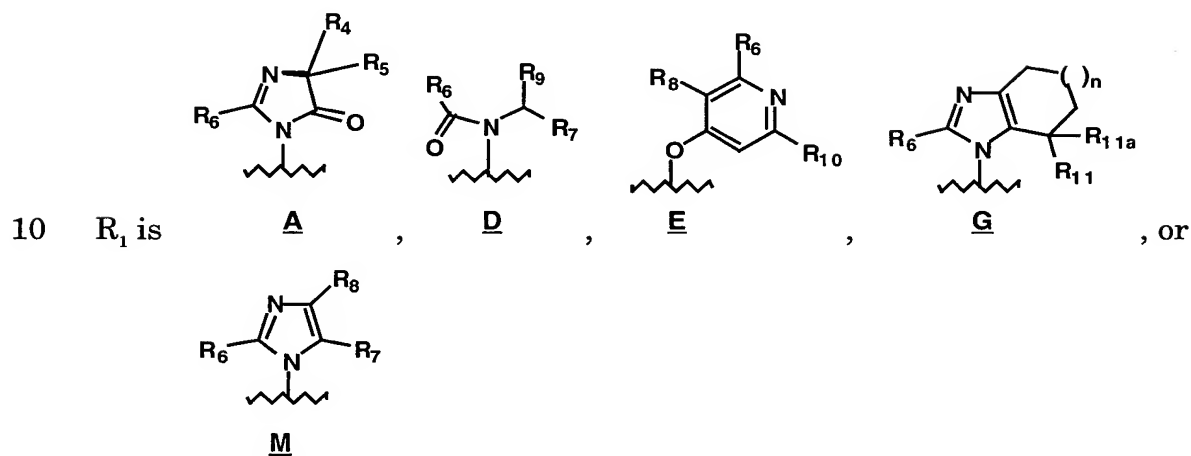
wherein

$R_1$ ,  $R_2$ ,  $R_{101}$  and  $R_{102}$  are as defined in claim 1; and

5  $R_{52}$  is chloro, bromo, iodo, or  $-\text{OSO}_2\text{CF}_3$ .

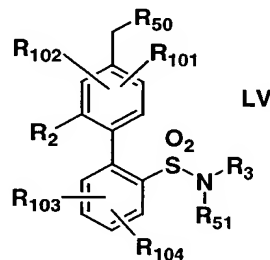
52. A compound of claim 51

wherein



53. A process for the preparation of a compound of claim 1, comprising at least one of the following steps:

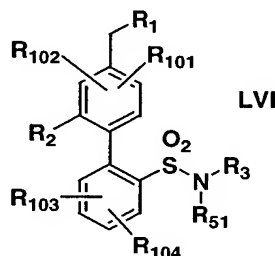
15 a) displacement of a leaving group  $R_{50}$  via the anion of a compound  $R_1\text{-H}$  from a compound of the formula



wherein  $R_1$ ,  $R_2$ ,  $R_3$ ,  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are as defined in claim 1,  $R_{50}$  is hydroxy, chloro, bromo, iodo,  $-\text{OSO}_2$ -alkyl or  $-\text{OSO}_2$ -aryl, and  $R_{51}$  is hydrogen or a suitable nitrogen protecting group, using a Mitsunobu reaction or  $S_N1$  or  $S_N2$  displacement reaction, with removal of said nitrogen

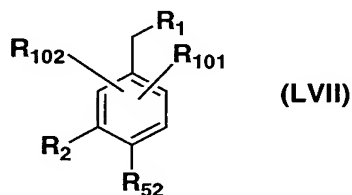
5 protecting group as appropriate;

b) removal of the nitrogen protecting group  $R_{51}$  from a compound of formula

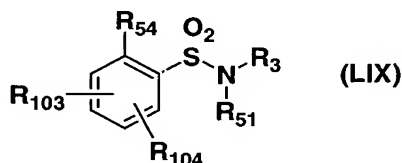


10 wherein  $R_1$ ,  $R_2$ ,  $R_3$ ,  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are as defined in claim 1, and  $R_{51}$  is a suitable nitrogen protecting group;

c) organometallic coupling of a compound of formula



15 with a compound of formula



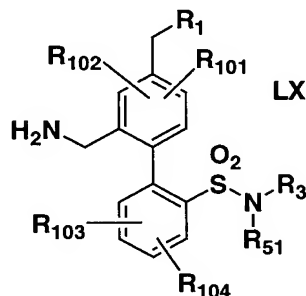
wherein

$R_1$ ,  $R_2$ ,  $R_3$ ,  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are as defined for a compound of formula I and  $R_{51}$  is a suitable nitrogen protecting group;

20  $R_{52}$  is chloro, bromo, iodo or  $-\text{OSO}_2\text{CF}_3$  and  $R_{54}$  is a boronic acid, boronic ester or stannane derivative;

or  $R_{52}$  is a boronic acid, boronic ester or stannane derivative and  $R_{54}$  is chloro, bromo, iodo or  $-\text{OSO}_2\text{CF}_3$ ;

d) acylation of a compound of the formula

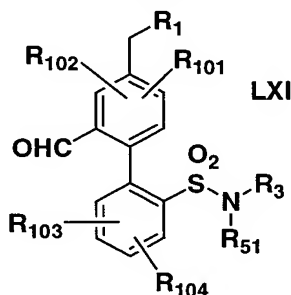


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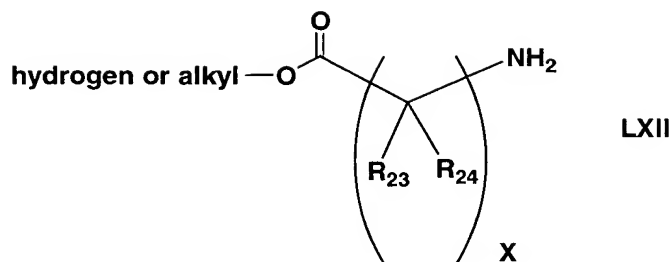
wherein  $R_1$ ,  $R_3$ ,  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are as defined in claim 1, and  $R_{51}$  is hydrogen or a suitable nitrogen protecting group, with an acylating agent of the formula  $R_{55}-(\text{C}=\text{O})R_{22}$ ,  $R_{19}\text{N}=\text{C}=\text{O}$ ,  $R_{55}-\text{CO}_2R_{18}$ ,  $R_{55}\text{SO}_2R_{22}$ , wherein  $R_{18}$ ,  $R_{19}$  and  $R_{22}$  are as defined in claim 1 and  $R_{55}$  is an activating group for an acid, or made using an acid activating agent, with removal of said nitrogen protecting group as appropriate; or

10

e) reductive amination of a compound of formula



15 wherein  $R_1$ ,  $R_3$ ,  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ , and  $R_{104}$  are as defined in claim 1, and  $R_{51}$  is hydrogen or a suitable nitrogen protecting group, with an amine of the formula



wherein  $R_{23}$ ,  $R_{24}$ , and  $x$  are as defined in claim 1, with removal of said nitrogen protecting group as appropriate.

5

54. A compound of claim 1, wherein said compound is selected from the group consisting of:

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethyl[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-ethyl[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)[1,1'-biphenyl]-2-sulfonamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethyl-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl[1,1'-biphenyl]-2-sulfonamide;

25 N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]- N-(3,4-dimethyl-5-isoxazolyl)-4'-  
 5 [[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl] [1,1'-biphenyl]-2-  
 10 sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]- N-(3,4-dimethyl-5-isoxazolyl)-4'-  
 15 [[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-4-isoxazolyl)-2'-ethyl-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-  
 20 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4'-  
 25 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-  
 30 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-4'-  
5 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-  
10 dimethyl-5-isoxazolyl)-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;

15 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;

20 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-  
25 sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-  
30 propyl-1-cycloheptimidazolyl)methyl]-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-

5 sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1,2,3,4-tetrahydrofuran-1-yl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,2,3,4-tetrahydrofuran-1-yl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,2,3,4-tetrahydrofuran-1-yl)[1,1'-

25 biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

30

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

5 N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

10 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

15 N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

20 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

25 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

30 4'-[[2-(2,2-Difluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;



4'-[[2-(2,2-Difluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

5 4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

10 4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

15 4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

20 4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

25 4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

30 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

5 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

15 1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

20 1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

25 1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

5 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

10 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

15 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

20 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

25 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;

30 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;

- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy)methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy)methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- 5 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy)methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy)methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- 10 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy)methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy)methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy)methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- 15 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy)methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy)methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy)methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- 20 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy)methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy)methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
- 25 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy)methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy)methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- 30 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy)methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;

5 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;

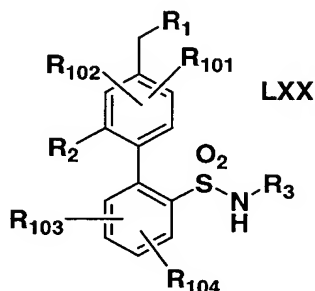
N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

10 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

and salts, enantiomers and diastereomers thereof.

55. A compound of formula LXX, or an enantiomer, diastereomer or salt thereof

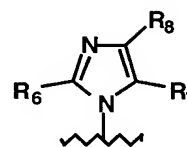
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wherein R<sub>2</sub>, R<sub>101</sub>, R<sub>102</sub>, R<sub>103</sub>, R<sub>104</sub> are as defined in claim 1;

R<sub>3</sub> is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen; and

20 R<sub>1</sub> is any group such that the resulting compound demonstrates affinity (IC<sub>50</sub>) for both the AT<sub>1</sub> receptor and ET<sub>A</sub> receptor of less than 5 micromolar at both receptors.



56. A compound of claim 5, wherein R<sub>1</sub> is

M

57. A compound of claim 56, wherein  $R_2$  is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and  $R_{101}$ ,  $R_{102}$ ,  $R_{103}$ ,  $R_{104}$  are each independently hydrogen, halogen, or alkyl.

5

58. A compound of claim 56, wherein  $R_2$  is  $-CH_2Y$ .

59. A compound of claim 58, wherein Y is Q.

60. A crystalline form of 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide having a melting point of about 148°C.

10

61. N2-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxobutyl)-L-valinamide or a salt, enantiomer or diastereomer thereof.

15

62. 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

20

63. 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2-fluoroethoxymethyl) [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

64. 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

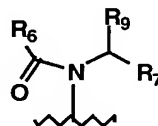
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65. N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethoxymethyl-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

30

66. N-(4,5-Dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

5 67. N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.



10 68. A compound of claim 5, where in R<sub>1</sub> is D.

69. A compound of claim 68, wherein R<sub>2</sub> is hydrogen, alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R<sub>101</sub>, R<sub>102</sub>, R<sub>103</sub>, R<sub>104</sub> are each independently hydrogen, halogen, or alkyl.

15 70. A compound of claim 68 wherein R<sub>2</sub> is -CH<sub>2</sub>Y.

71. A compound of claim 70, wherein Y is Q.

20 72. A method for the treatment of chronic obstructive pulmonary disease, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

25 73. A method for the alleviation of pain associated with prostate cancer, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

74. A method for preventing or reducing organ damage associated with the cell poliferative effects of endothelin, comprising the step of

administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

75. A method for reducing general morbidity and mortality  
5 associated with endothelin-dependent or angiotensin II-depedendent disorders, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

76. A method for the treatment of diabetic nephropathy, comprising  
10 the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

77. A method for the treatment of dementia, comprising the step of administering to a subject in need thereof an amount effective therefor of  
15 at least one compound of claim 1.

78. A compound of claim 1 wherein  $R_3$  is other than pyridyl when  $R_1$  is A.

79. The method of claim 26 wherein said compound of claim 1 is  
20 administered prior to, simultaneously with or following the administration of at least one ACE inhibitor.

80. The method of claim 79 wherein said ACE inhibitor is  
25 selected from captopril, zofenopril, fosinopril, ceranapril, alacepril, enalapril, delapril, pentopril, quinapril, ramipril, or lisinopril.

81. The method of claim 26 wherein said compound of claim 1 is  
30 administered prior to, simultaneously with or following the administration of at least one vasopepsidase inhibitor.



82. The method of claim 81 wherein said vasopepsidase inhibitor is selected from omapatrilat or gemopatrilat.

5 83. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one HMG CoA reductase inhibitor.

84. The method of claim 83 wherein said HMG CoA reductase inhibitor is selected from pravastatin, lovastatin, atorvastatin,  
10 simvastatin, NK-104 or ZD-4522.

85. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one anti-platelet agent.  
15

86. The method of claim 85 wherein said anti-platelet agent is selected from clopidigrel, ticlopidine, CS-747 or aspirin.

87. The method of claim 26 wherein said compound of claim 1 is  
20 administered prior to, simultaneously with or following the administration of at least one anti-diabetic agent.

88. The method of claim 87 wherein said anti-diabetic agent is selected from biguanides or biguanide/glyburide combinations.  
25

89. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one beta-adrenergic agent.

90. The method of claim 89 wherein said beta-adrenergic agent is selected from carvedilol or metoprolol.

5 91. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one mineralocorticoid receptor antagonist.

92. The method of claim 91 wherein said mineralocorticoid receptor antagonist is selected from spironolactone or eplerenone.  
10

93. The pharmaceutical composition of claim 46 further comprising at least one ACE inhibitor.

94. The pharmaceutical composition of claim 93 wherein said  
15 ACE inhibitor is selected from captopril, zofenopril, fosinopril, ceranapril, alacepril, enalapril, delapril, pentopril, quinapril, ramipril, or lisinopril.

95. The pharmaceutical composition of claim 46 further  
20 comprising at least one vasopepsidase inhibitor.

96. The pharmaceutical composition of claim 95 wherein said vasopepsidase inhibitor is selected from omapatrilat or gemopatrilat.

25 97. The pharmaceutical composition of claim 46 further comprising at least one HMG CoA reductase inhibitor.

98. The pharmaceutical composition of claim 97 wherein said HMG CoA reductase inhibitor is selected from pravastatin, lovastatin,  
30 atorvastatin, simvastatin, NK-104 or ZD-4522.

99. The pharmaceutical composition of claim 46 further comprising at least one anti-platelet agent.

5           100. The pharmaceutical composition of claim 99 wherein said anti-platelet agent is selected from clopidigrel, ticlopidine, CS-747 or aspirin.

10           101. The pharmaceutical composition of claim 46 further comprising at least one anti-diabetic agent.

15           102. The pharmaceutical composition of claim 101 wherein said anti-diabetic agent is selected from biguanides or biguanide/glyburide combinations.

            103. The pharmaceutical composition of claim 46 further comprising at least one beta-adrenergic agent.

20           104. The pharmaceutical composition of claim 103 wherein said beta-adrenergic agent is selected from carvedilol or metoprolol.

            105. The pharmaceutical composition of claim 46 further comprising at least one mineralocorticoid receptor antagonist.

25           106. The pharmaceutical composition of claim 105 wherein said mineralocorticoid receptor antagonist is selected from spironolactone or eplerenone.

107. The compound of claim 62 that is (+)-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide.

5            108. The compound of claim 62 that is (-)-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide.